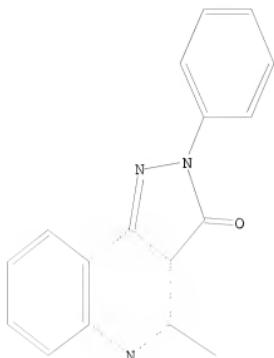


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G1 H,Ak

Structure attributes must be viewed using STN Express query preparation.

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PROJECTED ANSWERS:          2 TO       124
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FILE 'CAPLUS' ENTERED AT 15:50:14 ON 22 SEP 2008
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FILE COVERS 1907 - 22 Sep 2008 VOL 149 ISS 13
 FILE LAST UPDATED: 21 Sep 2008 (20080921/ED)

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L4  ANSWER 1 OF 2  CAPLUS  COPYRIGHT 2008 ACS on STN
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* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

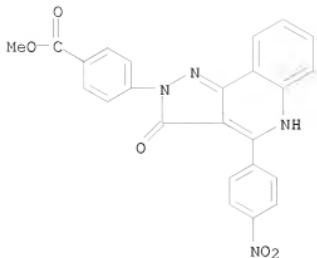
AB The invention relates to a preparation of novel pyrazolo[4,3-c]quinoline derivs. of formula I [wherein: Z is carboxylic acid or ester group; X is a bond or a divalent group selected from alkylene, or NHC(O), etc.; Y is O, S, N-oxide, or NH, etc.; R1 and R3 are independently selected from H, F, Cl, NO2, or CN, etc.; R2 (un)substituted C3-7cycloalkyl or phenyl] as immunomodulatory agents, useful for the treatment of rheumatoid arthritis, multiple sclerosis, diabetes, asthma, transplantation, systemic lupus erythematosis, and psoriasis. The title compds. are CD80 antagonists capable of inhibiting the interactions between CD80 and CD28. For instance, IC50 for the prepared pyrazoloquinoline derivative II was 3.4 μ M

(example 7).
 AN 2004:534202 CAPLUS
 DN 141:71540
 TI A preparation of pyrazolo[4,3-c]quinoline derivatives, useful as immunomodulatory agents
 IN Matthews, Ian Richard; Huxley, Philip; Magaraci, Filippo; Brennan, Chris James; Uddin, Muhammed Kamal; Pettersson, Lars Olof Goeran; Thriige, Dorthe Da Graca
 PA Active Biotech Ab, Swed.
 SO PCT Int. Appl., 28 pp.
 CODEN: PIXXD2

DT Patent
 LA English
 FAN.CNT 1

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PI	WO 2004055014	A1	20040701	WO 2003-SE1941	20031212
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OS MARPAT 141:71540
 IT 713141-32-3P
 RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (preparation of pyrazoloquinoline derivs. useful as immunomodulatory agents)
 RN 713141-32-3 CAPLUS
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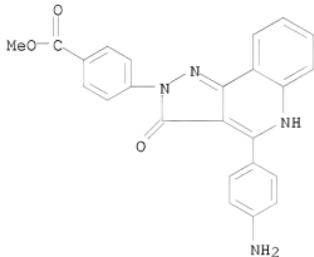
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 713141-39-0P 713141-40-3P 713141-41-4P
 713141-42-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of pyrazoloquinoline derivs. useful as immunomodulatory agents)

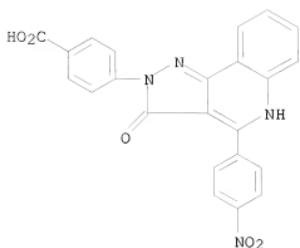
RN 713141-33-4 CAPLUS

CN Benzoic acid, 4-[4-(4-aminophenyl)-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]-, methyl ester (CA INDEX NAME)



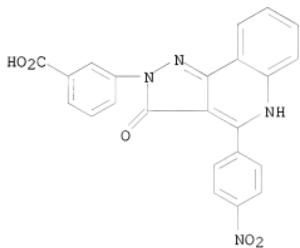
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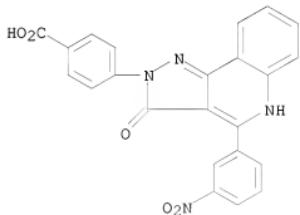
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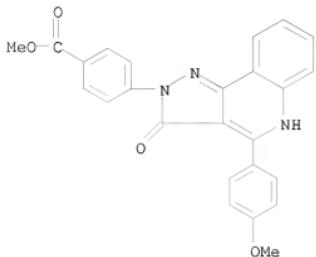


RN 713141-36-7 CAPLUS

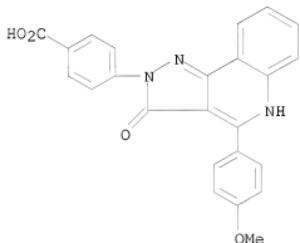
CN Benzoic acid, 4-[3,5-dihydro-4-(3-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)



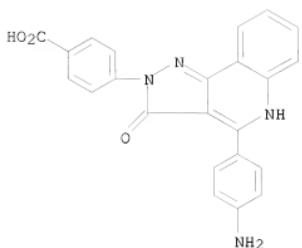
RN 713141-37-8 CAPLUS
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RN 713141-38-9 CAPLUS
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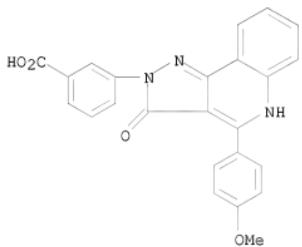


RN 713141-39-0 CAPLUS
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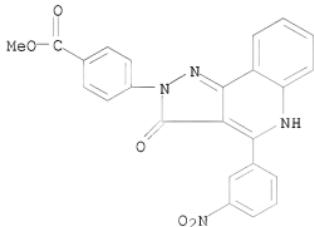
RN 713141-40-3 CAPLUS

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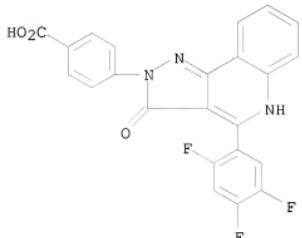


RN 713141-41-4 CAPLUS

CN Benzoic acid, 4-[3,5-dihydro-4-(3-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]-, methyl ester (CA INDEX NAME)



RN 713141-42-5 CAPLUS
 CN Benzoic acid, 4-[3,5-dihydro-3-oxo-4-(2,4,5-trifluorophenyl)-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)



RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN
 GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to novel heterocyclic compds., to methods for their preparation, to compns. containing them, and to methods and use for clin. treatment of medical conditions which may benefit from immunomodulation, including rheumatoid arthritis, multiple sclerosis, diabetes, asthma, transplantation, systemic lupus erythematosis, and psoriasis. More particularly, the invention relates to novel heterocyclic compds. I, which are CD80 antagonists capable of inhibiting the interactions between CD80 and CD28. In formula I, R1 and R3 independently represent H, F, Cl, Br, NO₂, CN, C1-C6 alkyl optionally substituted by F or Cl, or C1-C6 alkoxy optionally substituted by F; R2 represents H, or optionally substituted C1-C6 alkyl, C3-C7 cycloalkyl, or optionally substituted Ph; Y represents O, S, N-oxide, or N(R₅), wherein R₅ represents H or C1-C6 alkyl; X represents a bond or a divalent C1-C6 alkylene radical; R₄ represents -C(O)NR₆R₇, -NR₇C(O)R₆, -NR₇C(O)OR₆, -NHC(O)NHR₆, or -NHC(S)NHR₆, wherein R₆ represents H, or a radical of formula -(Alk)_b-Q wherein b = 0-1 and Alk is an optionally substituted divalent straight chain or branched C1-C12 alkylene, C2-C12 alkenylene or C2-C12 alkynylene radical which may be interrupted by one or more non-adjacent -O-, -S- or -N(R₈)- radicals wherein R₈ represents H or C1-C4 alkyl, C3-C4 alkenyl, C3-C4 alkynyl, or C3-C6 cycloalkyl, and Q represents H, CF₃, OH, SH, NR₈R₈ wherein each R₈ may be the same or different, an ester group, or an optionally substituted Ph, C3-C7 cycloalkyl, C5-C7 cycloalkenyl or heterocyclic ring having from 5 to 8 ring atoms; and R₇ represents H or C1-C6 alkyl; or when taken

together with the atom or atoms to which they are attached, R6 and R7 form an optionally substituted heterocyclic ring having from 5 to 8 ring atoms. Approx. 170 example compds. and several intermediates were prepared. For instance, invention compound II (claimed individually) was prepared in 5 steps: (1) cyclocondensation of 3-cyclopropyl-3-oxopropionic acid Me ester with Et 2-aminobenzoate to give a quinolone derivative, (2) conversion of the quinolone ester to a chloroquinolone ester with POC13, (3) cyclocondensation of the latter with 4-hydrazinobenzoic acid to form the pyrazole ring, (4) conversion of the free acid group to an acid chloride, and (5) amidation with H2N(CH2)3NMe2. In a cell-free, Eu/APC-based, homogeneous time-resolved fluorescence (HTRF) assay, used to determine inhibition of CD80-CD28 interaction, II had EC50 < 1 μ M.

AN 2004:467892 CAPLUS

DN 141:38606

TI Pyrazoloquinolines and analogs with CD80 antagonist immunomodulating activity, and their preparation, pharmaceutical compositions, and use
IN Matthews, Ian Richard; Coulter, Thomas Stephen; Ghironi, Chiara; Brennan, Chris James; Uddin, Muhammed Kamal; Pettersson, Lars Olof Goeran; Da Graca Thriige, Dorthe; Huxley, Philip

PA Active Biotech AB, Swed.

SO PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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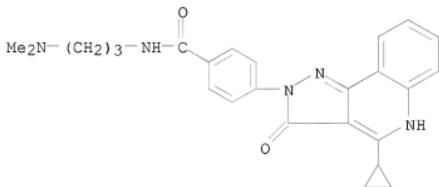
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 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of pyrazoloquinolines and analogs as CD80 antagonists and immunomodulators)

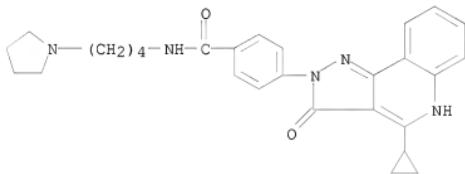
RN 702704-93-6 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[3-(dimethylamino)propyl]- (CA INDEX NAME)



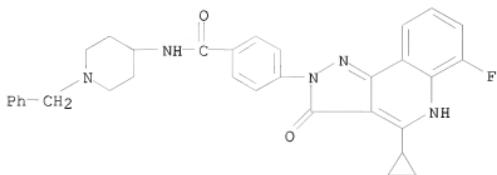
RN 702705-30-4 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[4-(1-pyrrolidinyl)butyl]- (CA INDEX NAME)



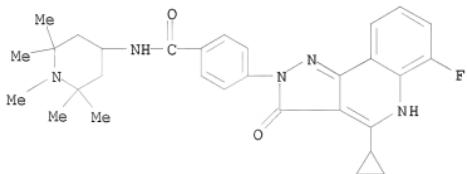
RN 702705-51-9 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[1-(phenylmethyl)-4-piperidinyl]- (CA INDEX NAME)



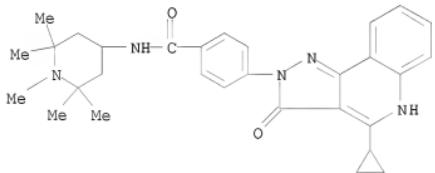
RN 702705-52-0 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(1,2,2,6,6-pentamethyl-4-piperidinyl)- (CA INDEX NAME)



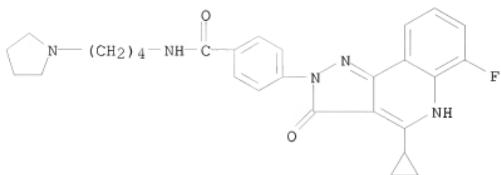
RN 702705-55-3 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(1,2,2,6,6-pentamethyl-4-piperidinyl)- (CA INDEX NAME)

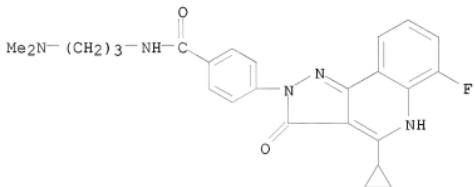


RN 702705-56-4 CAPLUS

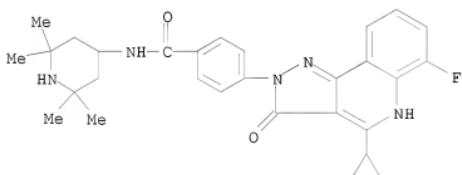
CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[4-(1-pyrrolidinyl)butyl]- (CA INDEX NAME)



RN 702705-58-6 CAPLUS
 CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(3-(dimethylamino)propyl)- (CA INDEX NAME)

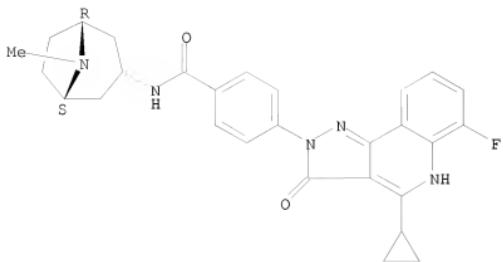


RN 702705-59-7 CAPLUS
 CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(2,2,6,6-tetramethyl-4-piperidinyl)- (CA INDEX NAME)

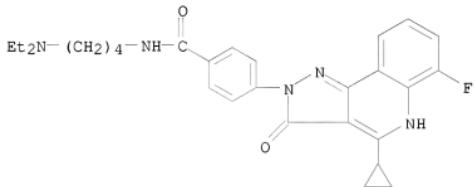


RN 702705-60-0 CAPLUS
 CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[(3-endo)-8-methyl-8-azabicyclo[3.2.1]oct-3-yl]- (CA INDEX NAME)

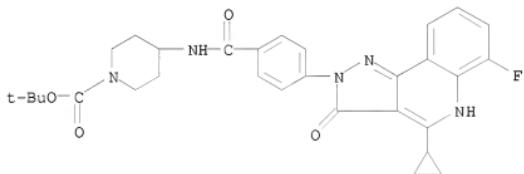
Relative stereochemistry.



RN 702705-61-1 CAPLUS
 CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[4-(diethylamino)butyl]- (CA INDEX NAME)

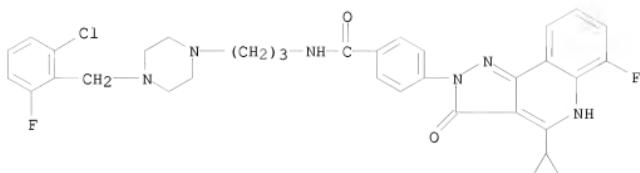


RN 702705-62-2 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)benzoyl]amino-, 1,1-dimethylethyl ester (CA INDEX NAME)



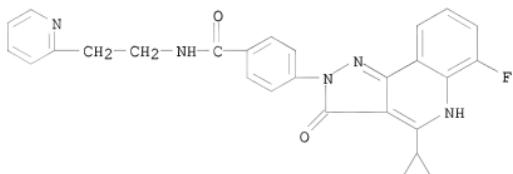
RN 702705-63-3 CAPLUS
 CN Benzamide, N-[3-[(2-chloro-6-fluorophenyl)methyl]-1-piperazinyl]propyl]-4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-

y1)- (CA INDEX NAME)



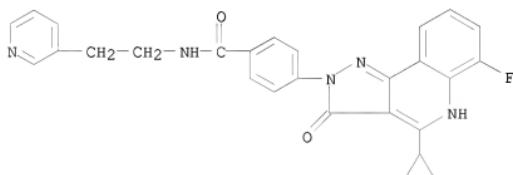
RN 702705-64-4 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(2-(2-pyridinyl)ethyl)- (CA INDEX NAME)



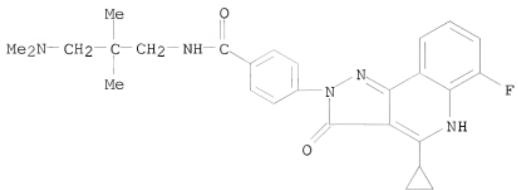
RN 702705-65-5 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(2-(3-pyridinyl)ethyl)- (CA INDEX NAME)



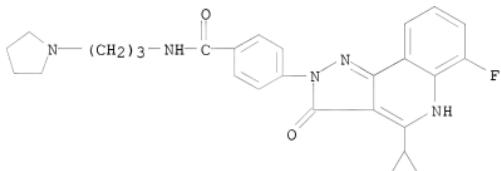
RN 702705-66-6 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[3-(dimethylamino)-2,2-dimethylpropyl]- (CA INDEX NAME)



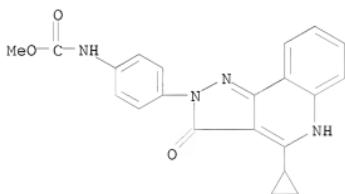
RN 702705-67-7 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[3-(1-pyrrolidinyl)propyl]- (CA INDEX NAME)



RN 702705-78-0 CAPLUS

CN Carbamic acid, [4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)phenyl]-, methyl ester (9CI) (CA INDEX NAME)



IT 702706-05-6P, 4-(4-Cyclopropyl-3-oxo-3,5-dihydropyrazolo[4,3-c]quinolin-2-yl)benzoic acid 702706-06-7P, 4-(4-Cyclopropyl-3-

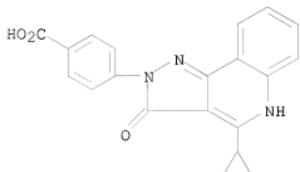
oxo-3,5-dihydropyrazolo[4,3-c]quinolin-2-yl)benzoyl chloride

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(intermediate; preparation of pyrazoloquinolines and analogs as CD80 antagonists and immunomodulators)

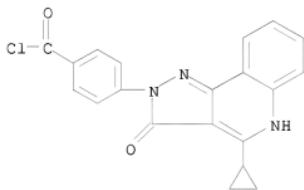
RN 702706-05-6 CAPLUS

CN Benzoic acid, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)- (CA INDEX NAME)



RN 702706-06-7 CAPLUS

CN Benzoyl chloride, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)- (CA INDEX NAME)



RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT